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MDTKHFLPLDFSTQVNSSLTSPTGRGSMAPSLHPSLGP GIGSPGQLHSPISTLSSPINGMG
PPFSVISSPMGPHSMSVPTTPTLGFSTGSPQLSSPMNPVSSSEDIKPPLGLNGVLKVPAPHS
GNMASFTKHICAICGDRSSGKH YGVYSCEGCKGFFKRTVRKDLTYTCRDNKDCLIDKRQRNR
CQYCRYQKCLAMGMKREAVQEERQRGKDRNENEVESTSSANEDMPVERILEAELAVEPKTET
YVEANMGLNPSSPNDPVTNICQAAADKQLFTLVEWAKRIPHFSELPLDDQVILLRAGWNELLI
ASFHSRISIAVKDGILLATGLHVHRNSAHSAGVGAI FDRVLTEL VSKMRDMQMDKTELGCLRA
IVLFNPDSKGLSNPAEVEALREKVYASLEAYCKHKYPEQPGRF AKLLLRLPALRSIGLKCLE
HLFFFKLIGDTPIDTFLMEMLEAPHQMT

FIGURE 1A

MSWAARPPFLPQRHAAGQCGPVGVRKEMHCGVASRWRRRRPWLDPAAAAAAAVAGGEQQTPE
PEPGEAGRDGMGDSGRDSRSPDSSSPNLPQGVPPPSPPGPPLPPSTAPSLGGSGAPPPPPM
PPPPLGSPFPVISSSMGSPGLPPPAPPFGSPVSSPQINSTVSLPGGGSGPPEDVKPPVLGV
RGLHCPPPPGGPGAGKRLCAICGDRSSGKH YGVYSCEGCKGFFKRTIRKDLTYSCRDNKDCT
VDKRQRNRCQYCRYQKCLATGMKREAVQEERQRGKDKDGDGEGAGGAPEEMPVDRILEAELA
VEQKSDQGVVEGPGGTGGSGSSPNDPVTNICQAAADKQLFTLVEWAKRIPHFSSSLPLDDQVILL
RAGWNELLIASFHSRISIDVRDGILLATGLHVHRNSAHSAGVGAI FDRVLTEL VSKMRDMRMD
KTELGCLRAIILFNPDAKGLSNPSEVEVLREKVYASLETYCKQKYPEQQGRFAKLLLRLPAL
RSIGLKCLEHLFFFKLIGDTPIDTFLMEMLEAPHQLA

FIGURE 1B

MYGNYSHFMKFPAGYGGSPGHTGSTSMSPSAALSTGKPMDSHPSYTDTPVSAPRTL SAVGTP
LNALGSPYRVITSAMGPPSGALAAPPGINLVAPPSSQLNVVNSVSSSEDIKPLPGLPGIGNM
NYPSTSPGSLVKHICAICGDRSSGKH YGVYSCEGCKGFFKRTIRKDLIYTCRDNKDCLIDKR
QRNRCQYCRYQKCLVMGMKREAVQEERQSRERA ESEAECATSGHEDMPVERILEAELAVEP
KTESYGDMMNENSTNDPVTNICHAADKQLFTLVEWAKRIPHFSDLTLEDQVILLRAGWNELL
IASFSHRSVSVQDGILLATGLHVHRSSAHSAGVGSIFDRVLTEL VSKMKDMQMDKSELGCLR
AIVLFNPDAKGLSNPSEVETLREKVYATLEAYTKQKYPEQPGRF AKLLLRLPALRSIGLKCL
EHLFFFKLIGDTPIDTFLMEMLETPLQIT

FIGURE 1C

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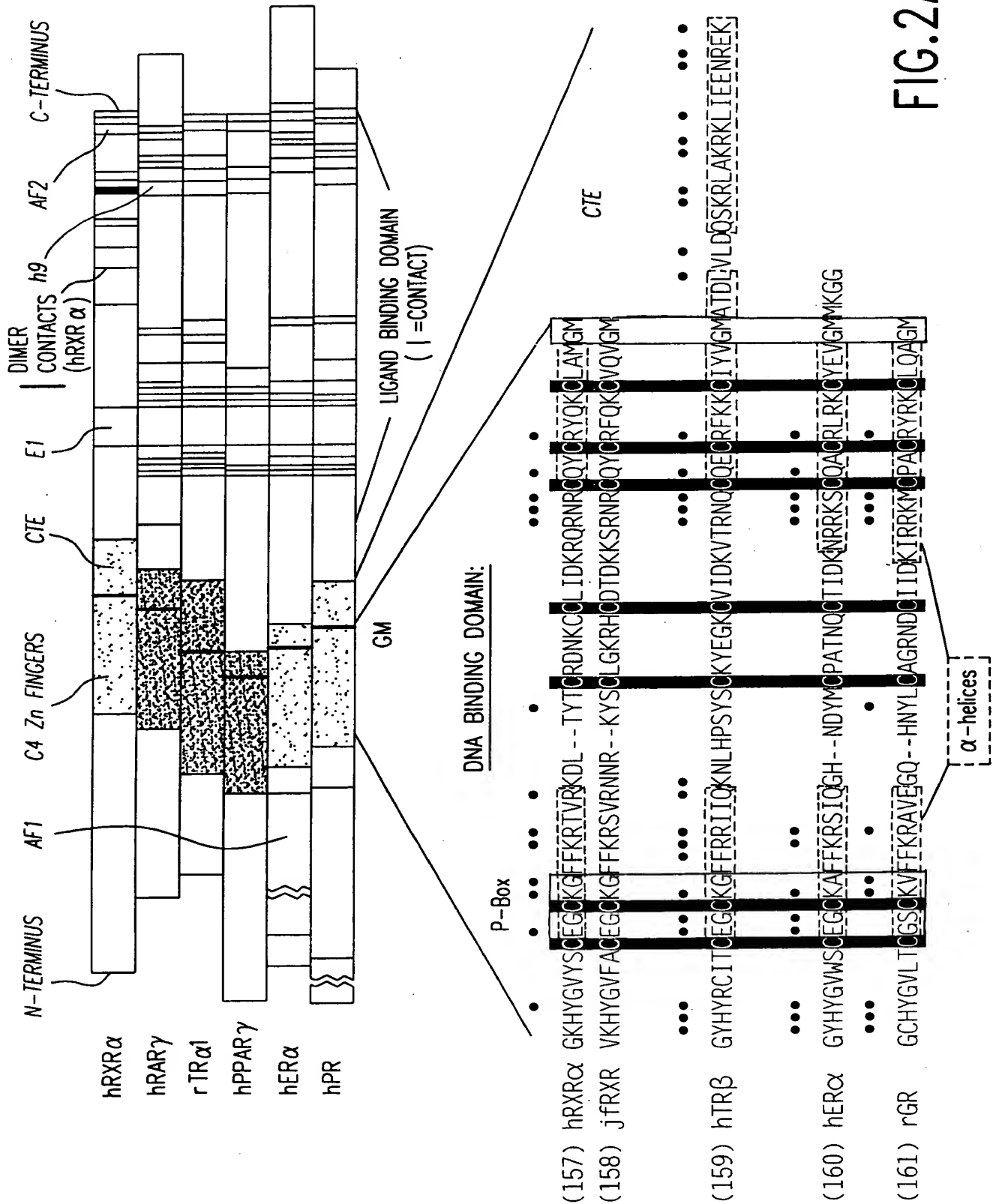


FIG. 2A

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E1 Region		Heptad 9 (h9)		AF2 Domain	
		ϕ	ϕ	ϕ	ϕ
(162) hRXR α	WAKRIPHFSELPDQVILL	(172) hRXR α	LLRLPALR	(182) hRXR α	FLMEML
(163) jFRXR	WAKRLPHFRDLSIADQVLL	(173) jFRXR	VILRIPALR	(183) jFRXR	FLLDML
(164) hTR α 1	FAKKLPMFSELPCEQIILL	(174) hTR α 1	LLMKVTDLR	(184) hTR α 1	LFLEVF
(165) hRAR γ	FAKRLPGFTGLSIADQITLL	(175) hRAR γ	MLMKITDLR	(185) hRAR γ	LTREML
(166) hPPAR γ	YAKSIPGFWNLNDQVTLL	(176) hPPAR γ	LLQKMTDLR	(186) hPPAR γ	LLQEIV
(167) hLXR	FAKQLPGFLQLSREDQIALL	(177) hLXR	MLMKLVSLR	(187) hLXR	LLSEIW
(168) hVDR	FAKMIPGFRDLTSEDQIVLL	(178) hVDR	MTQKLADLR	(188) hVDR	LVLEVF
(169) hER	WAKRVPGFVDLTLDQVHLL	(179) hER	LLLILSHIR	(189) hER	LLLEML
(170) hGR	WAKAIPGFRNLHLDQMTLL	(180) hGR	LTKLLDSMH	(190) hGR	MLAEII
(171) hPR	WSKSLPGFRNLHIDDQITLI	(181) hPR	LTKLLDNLH	(191) hPR	MMSEVI
α -helix 3		α -helix 10/11		α -helix 12	

FIG.2B

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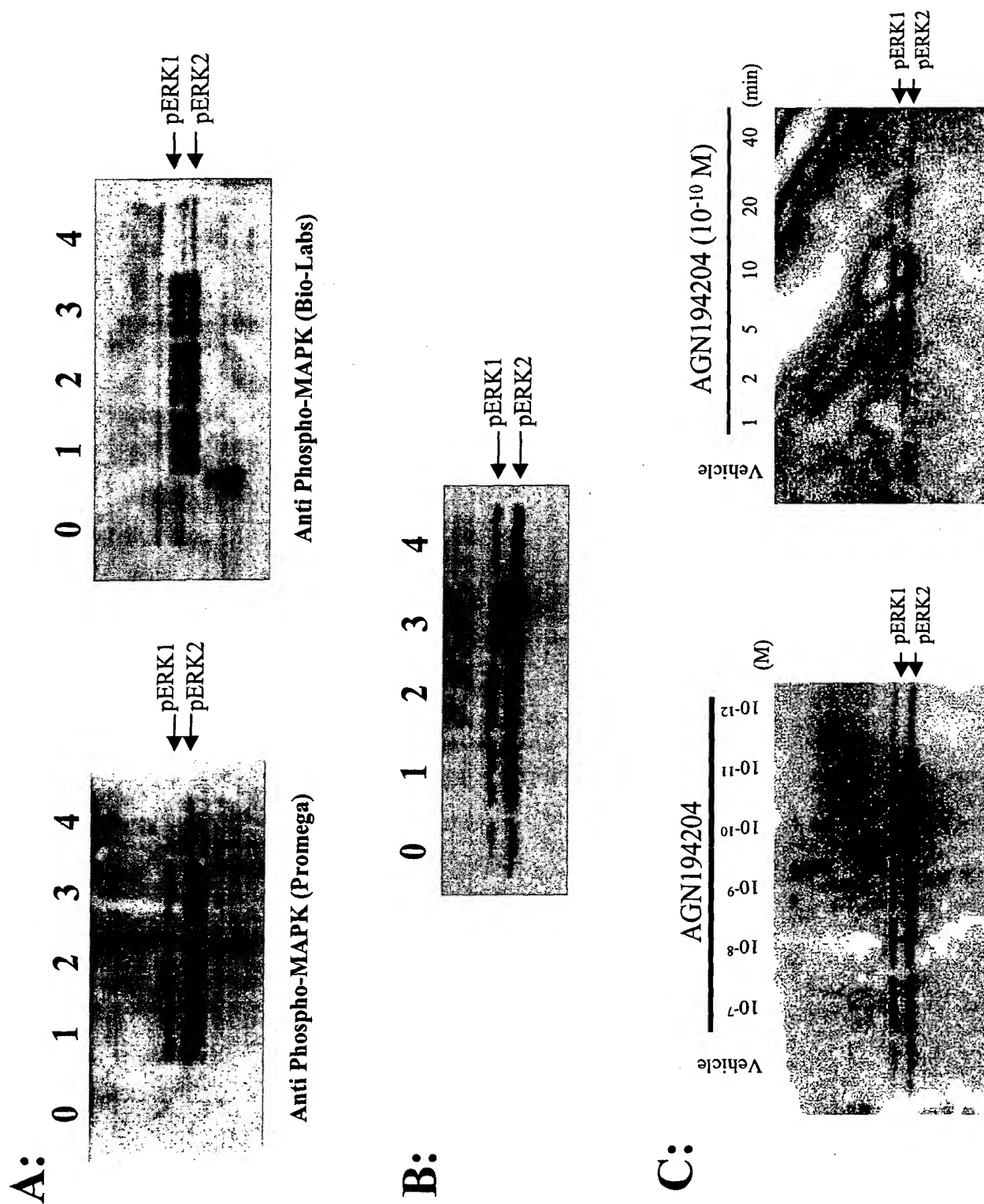


FIGURE 3

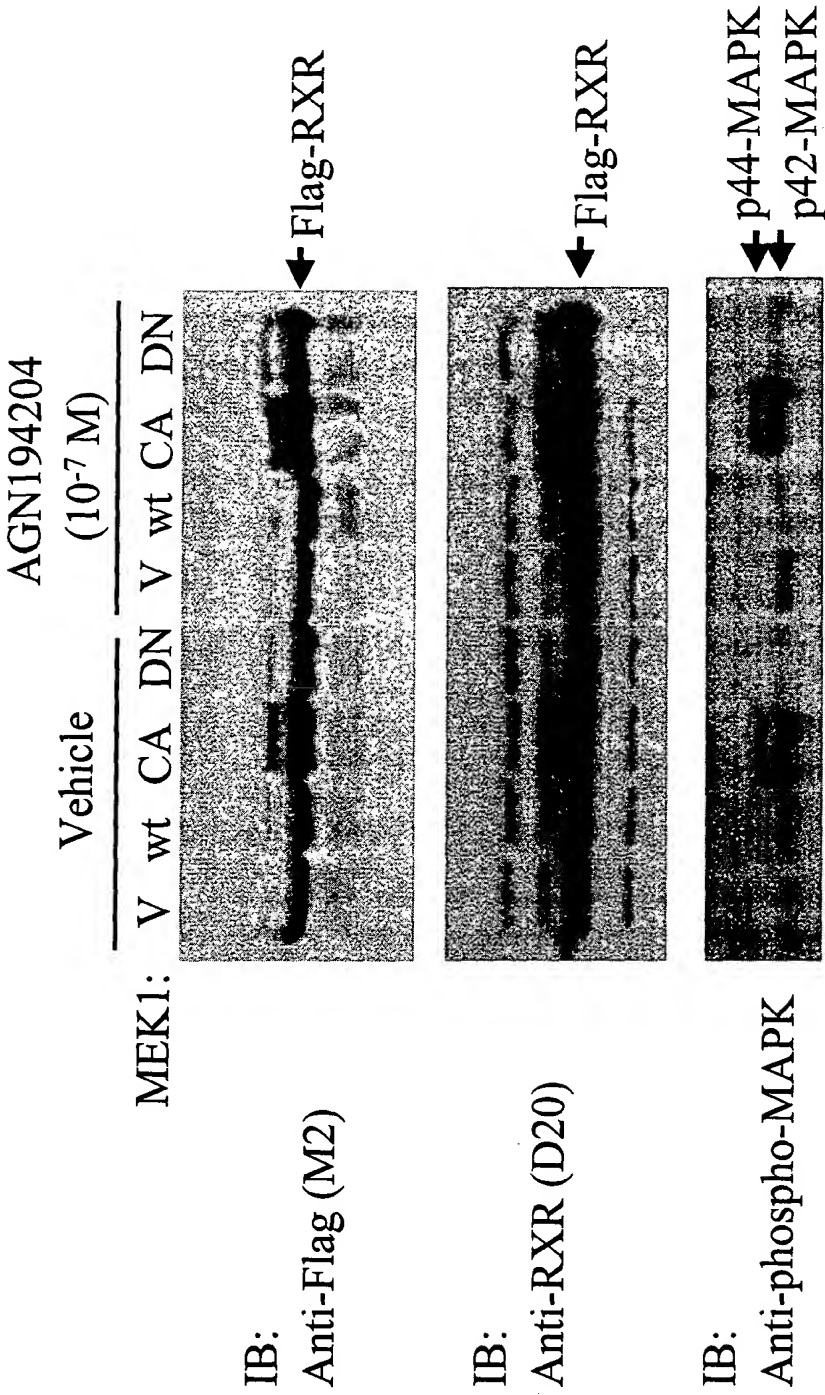


FIGURE 4

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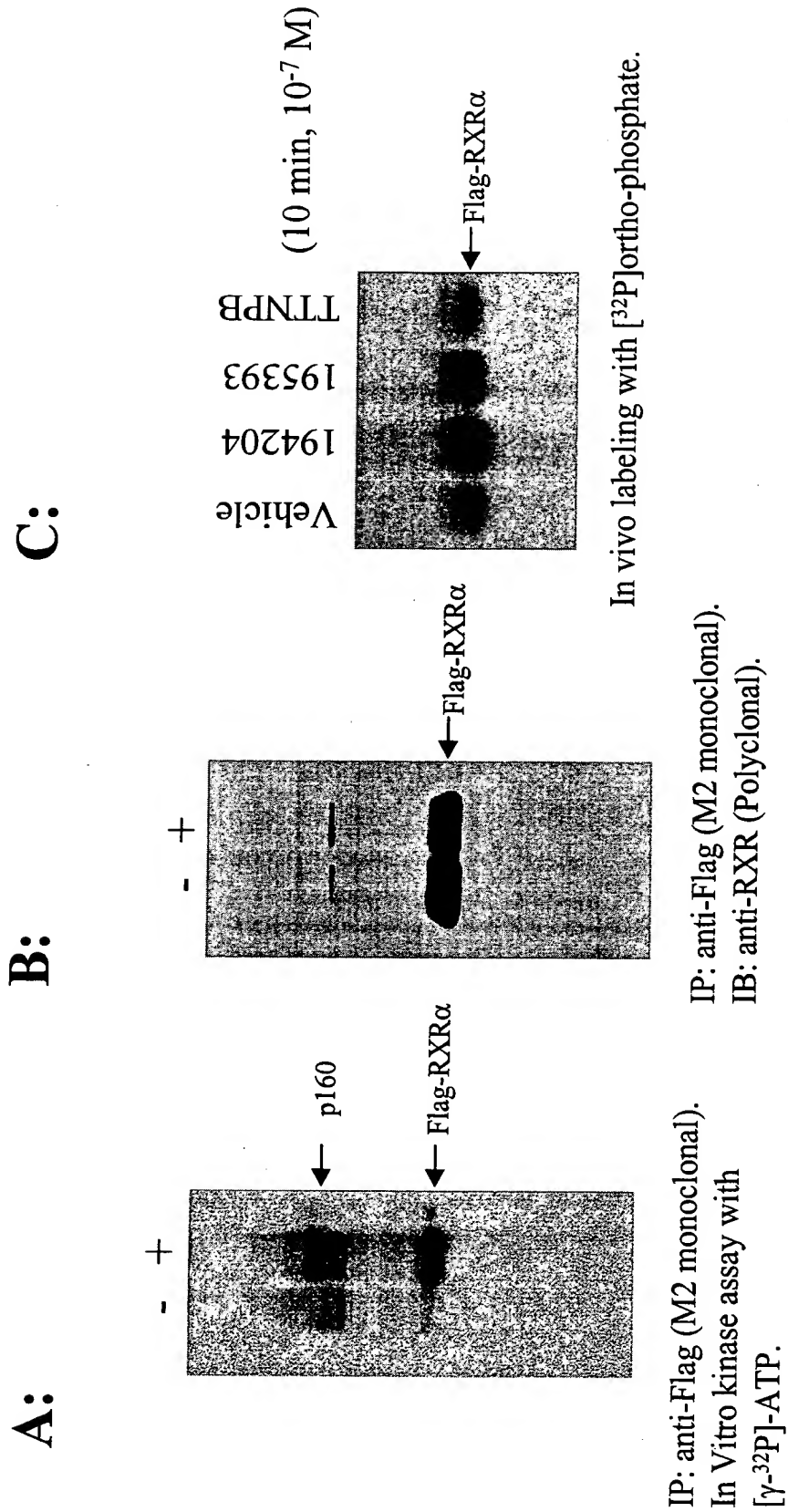
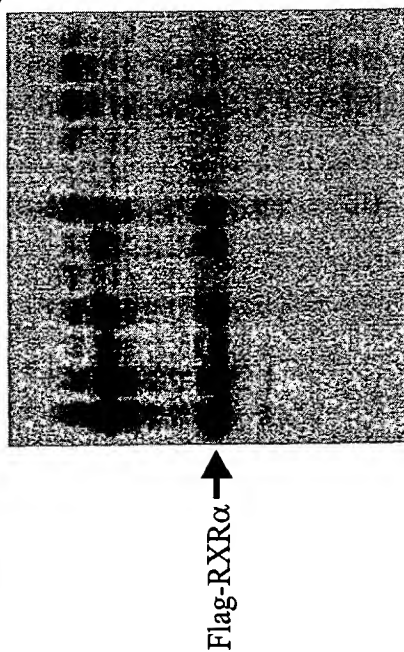


FIGURE 5

\ddot{A}

IP Buffer: - + - + - + - + - + - + AGN194204

Stimulation: 10⁻⁶ 10⁻⁷ 10⁻⁸ 10⁻⁹ 10⁻¹⁰ 10⁻¹¹ (M) AGN194204 (10 min)



Western blot analysis showing the levels of Flag-RXR and p160 in various cell lines. The lanes are labeled: Vector, AGN194204, Vehicle, AGN195029, AGN192620, AGN195023, AGN195393, and AGN195184. The top row of bands is labeled p160, and the bottom row is labeled Flag-RXR. The Flag-RXR bands are present in all lanes except the Vector lane, indicating successful transfection. The p160 bands are present in all lanes, indicating the presence of the endogenous protein.

FIGURE 6

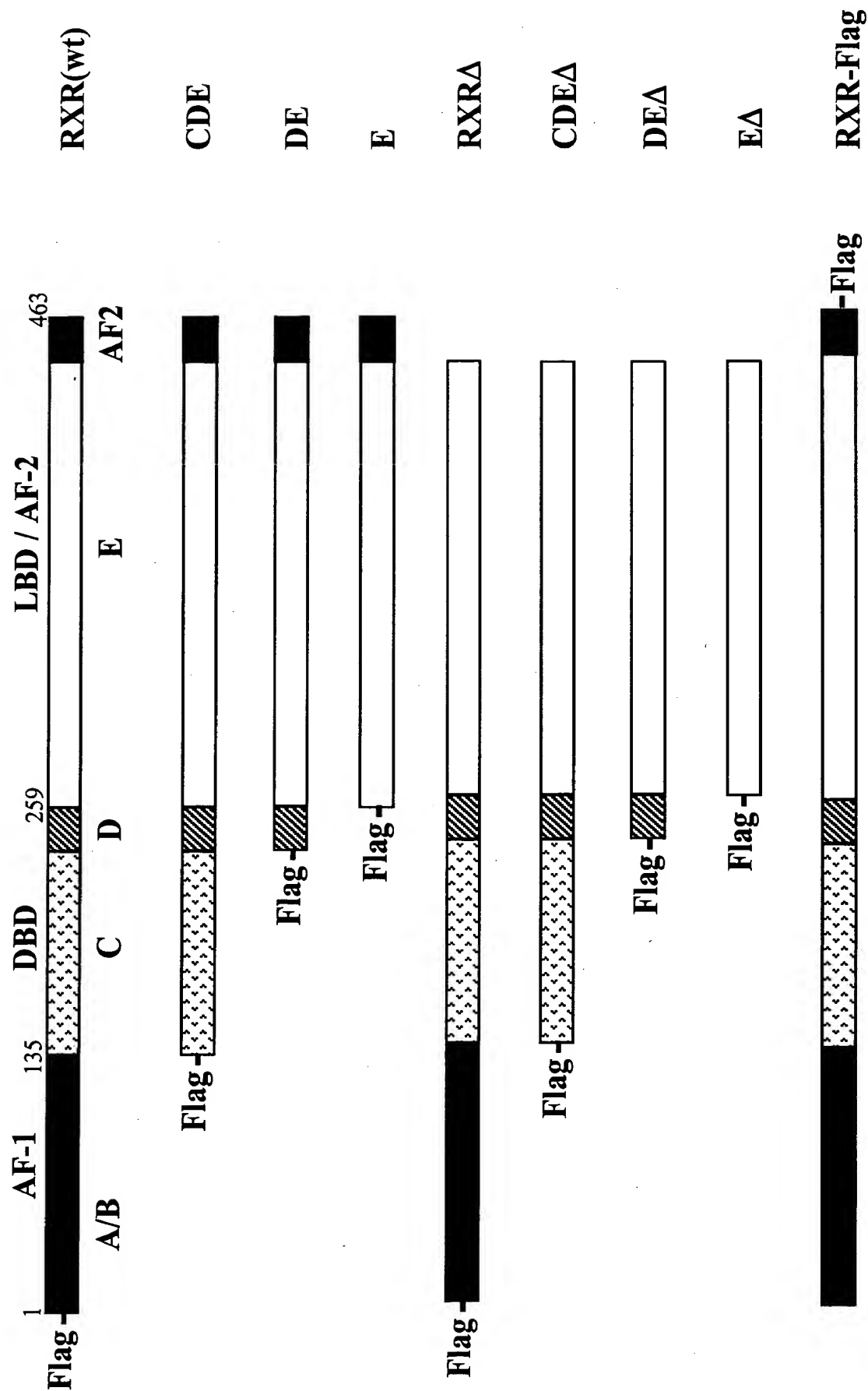
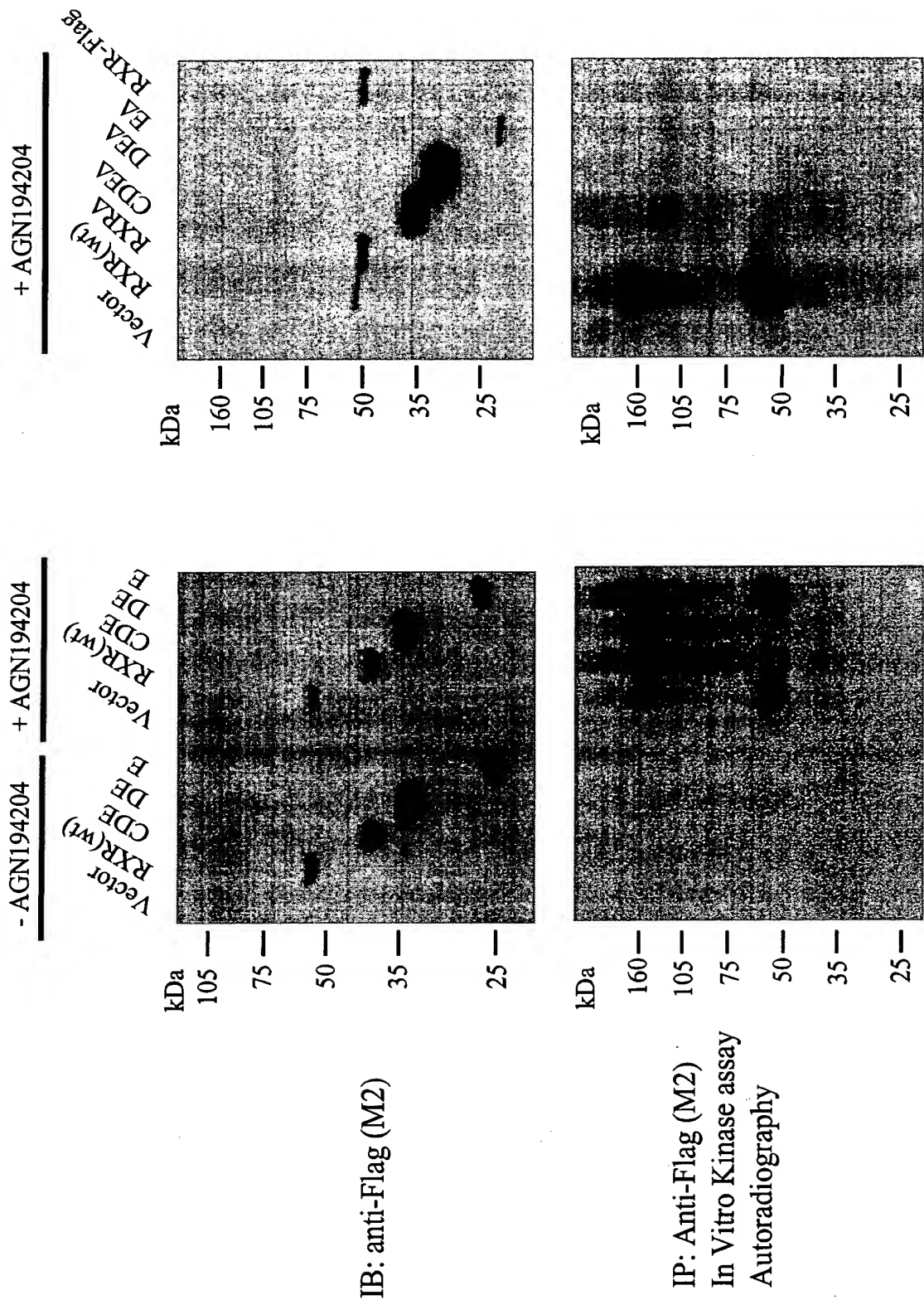


FIGURE 7

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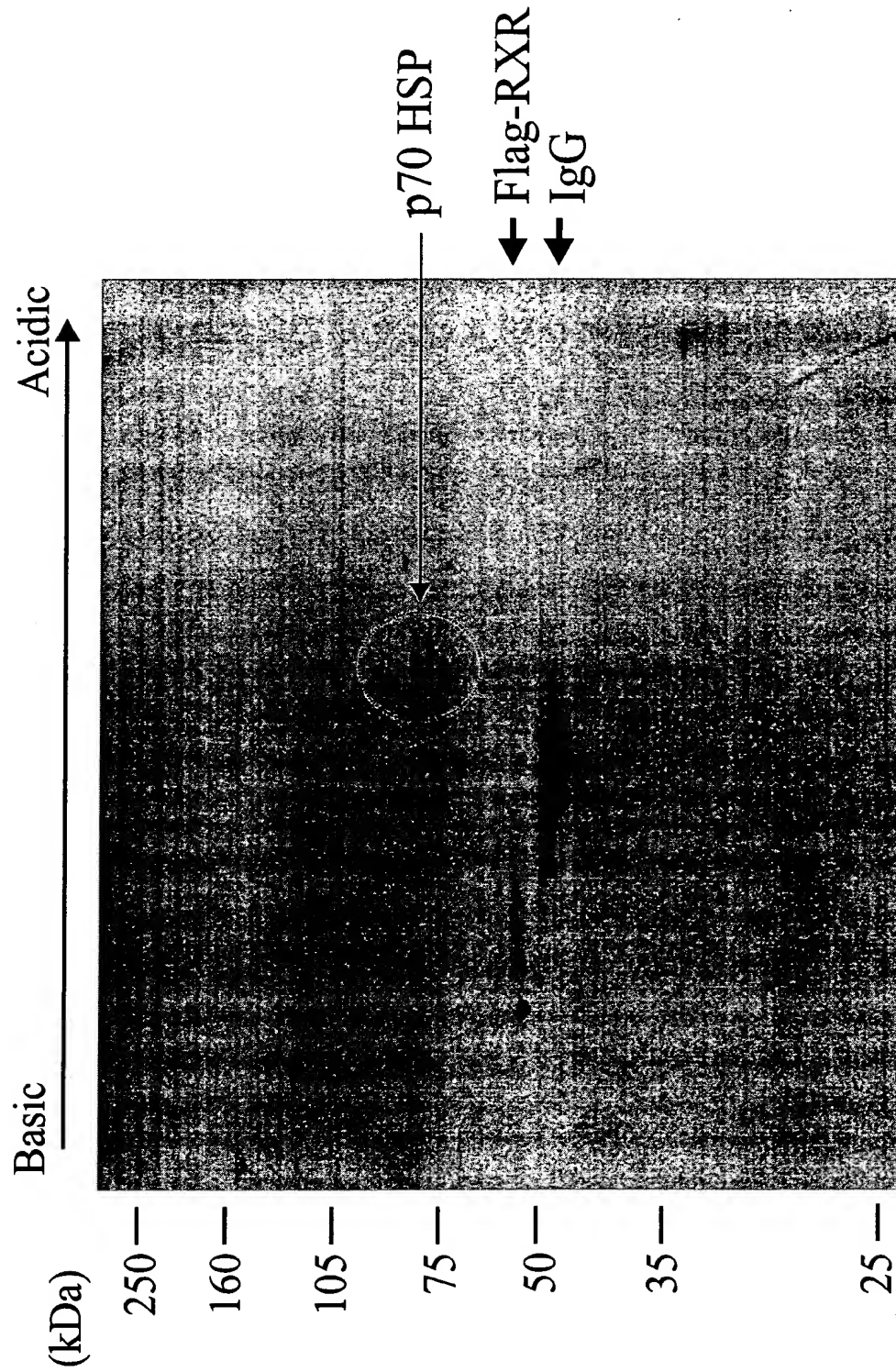


FIGURE 9

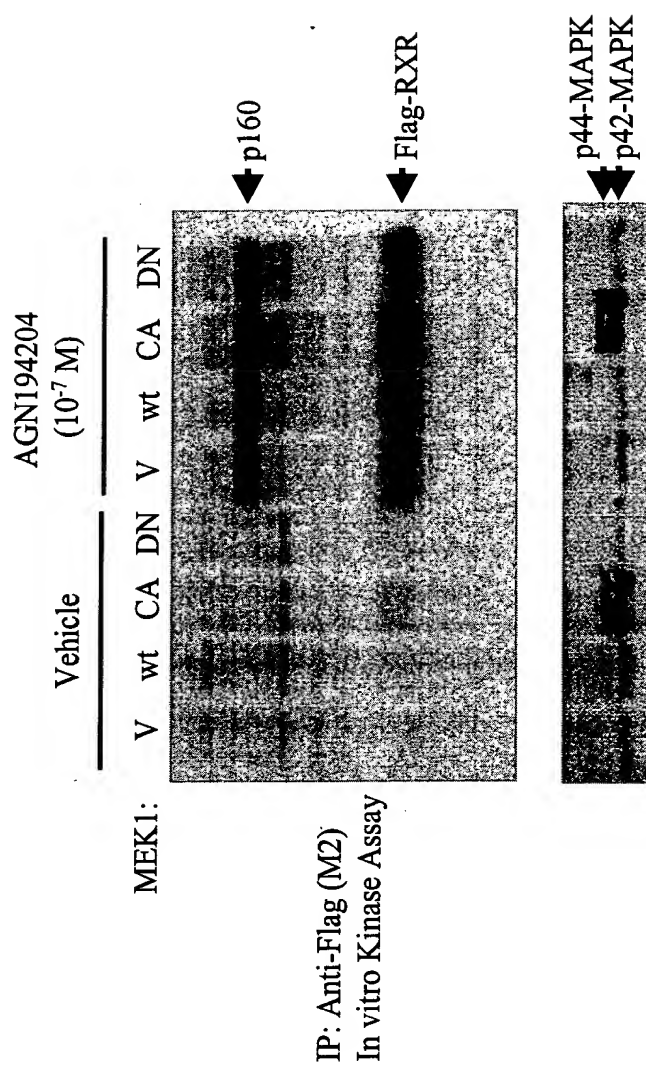
\ddot{A} 

FIGURE 11

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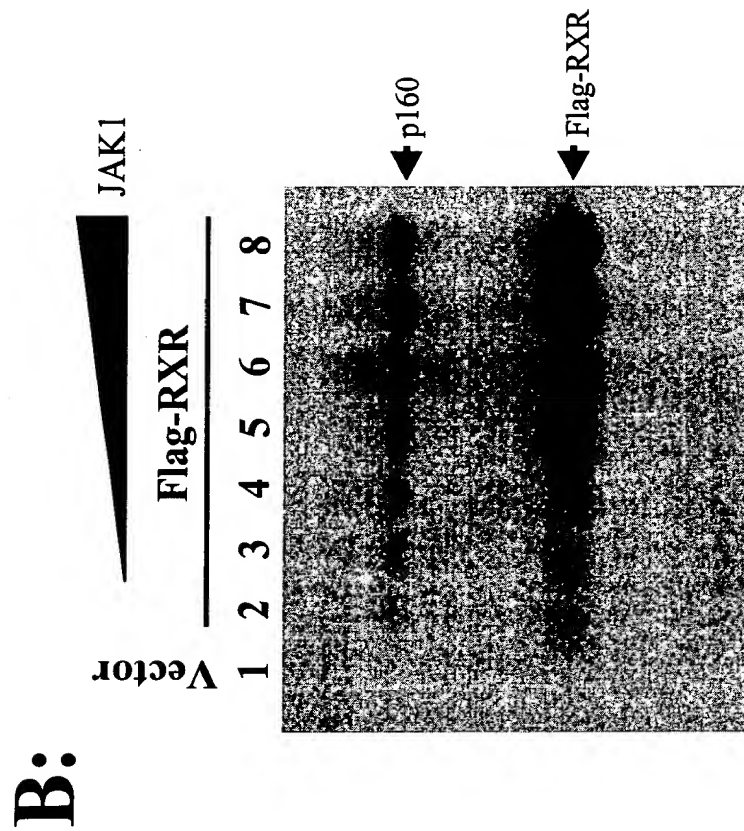


FIGURE 11 CONTINUED

Figure 1 is a plot of Counts versus Concentration [M]. The y-axis is labeled 'Counts' and ranges from 0 to 2,500,000. The x-axis is labeled 'Concentration [M]' and ranges from -9 to -5. The plot shows data points with error bars for three AGN sources: AGN 194204 (squares), AGN 195393 (triangles), and AGN 195741 (inverted triangles). Two sigmoidal curves are fitted to the data, showing a sharp increase in counts between -8 M and -6 M. The data points for AGN 194204 and AGN 195741 are clustered around the curves, while AGN 195393 data points are significantly lower, near the baseline.

Concentration [M]	AGN 194204 (Counts)	AGN 195393 (Counts)	AGN 195741 (Counts)
-9	~100,000	~100,000	~100,000
-8	~200,000	~100,000	~200,000
-7	~1,200,000	~100,000	~800,000
-6	~2,200,000	~100,000	~2,200,000
-5	~2,200,000	~100,000	~2,200,000

	AGN 194204	AGN 195393	AGN 195741
EC50	7.7150e-008		1.7540e-007

Figure 1 is a plot of Counts versus Concentration [M]. The y-axis represents Counts, ranging from 0 to 5,000,000. The x-axis represents Concentration [M], ranging from -9 to -5. Data points with error bars are shown for three AGN sources: AGN 194204 (squares), AGN 195393 (triangles), and AGN 195741 (inverted triangles). Two sigmoidal curves are fitted to the data, with the upper curve corresponding to AGN 194204 and the lower curve to AGN 195393. AGN 195741 data points are near the baseline.

Concentration [M]	AGN 194204 (Counts)	AGN 195393 (Counts)	AGN 195741 (Counts)
-9	~100,000	~100,000	~100,000
-8	~200,000	~100,000	~100,000
-7	~1,500,000	~1,200,000	~1,000,000
-6	~3,500,000	~2,800,000	~100,000
-5	~4,500,000	~3,200,000	~100,000

	AGN 194204	AGN 195393	AGN 195741
EC50	1.2010e-007		2.2510e-007

FIGURE 12

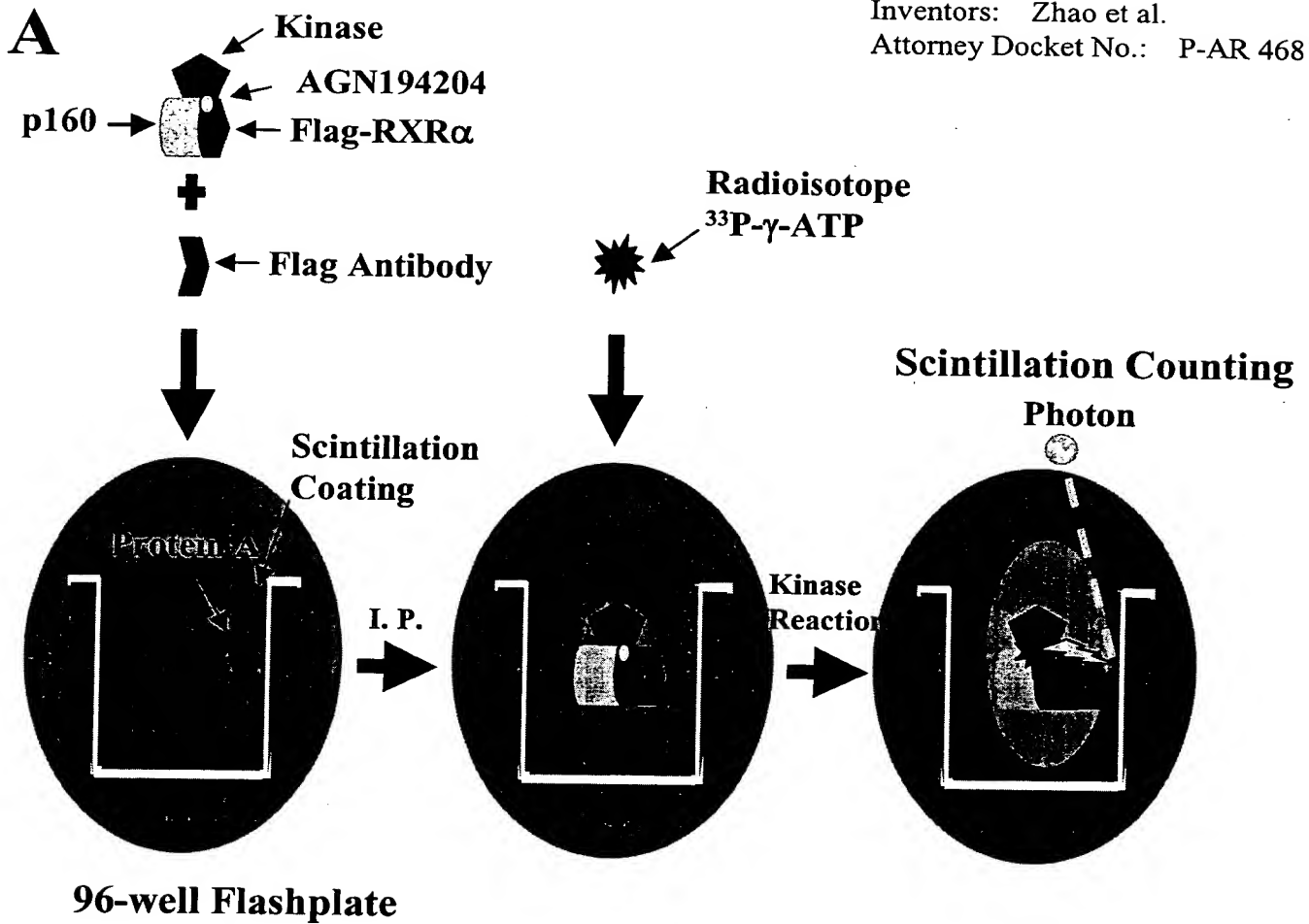
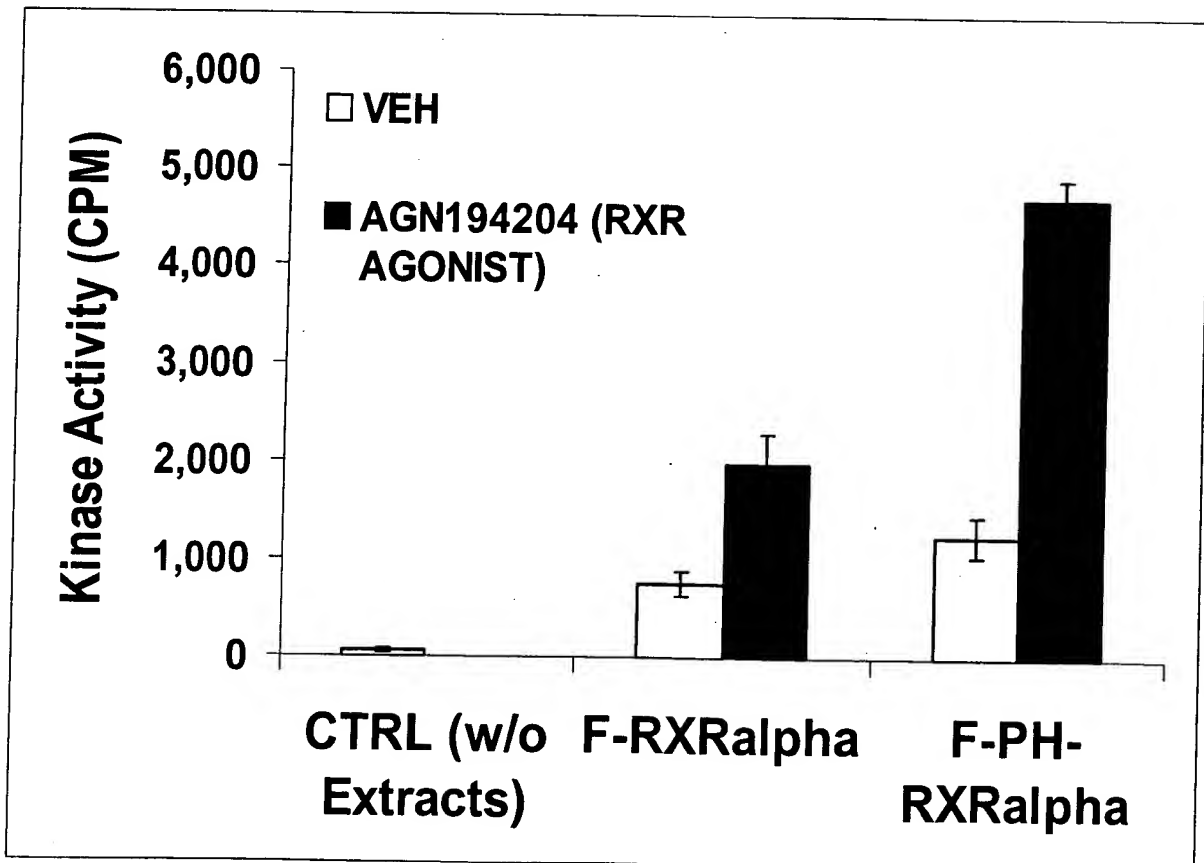
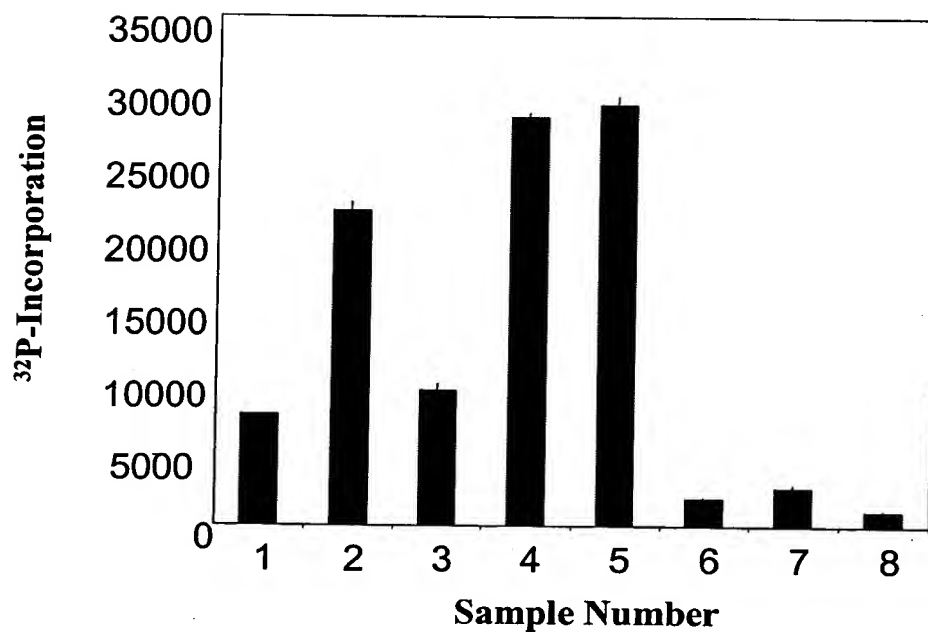
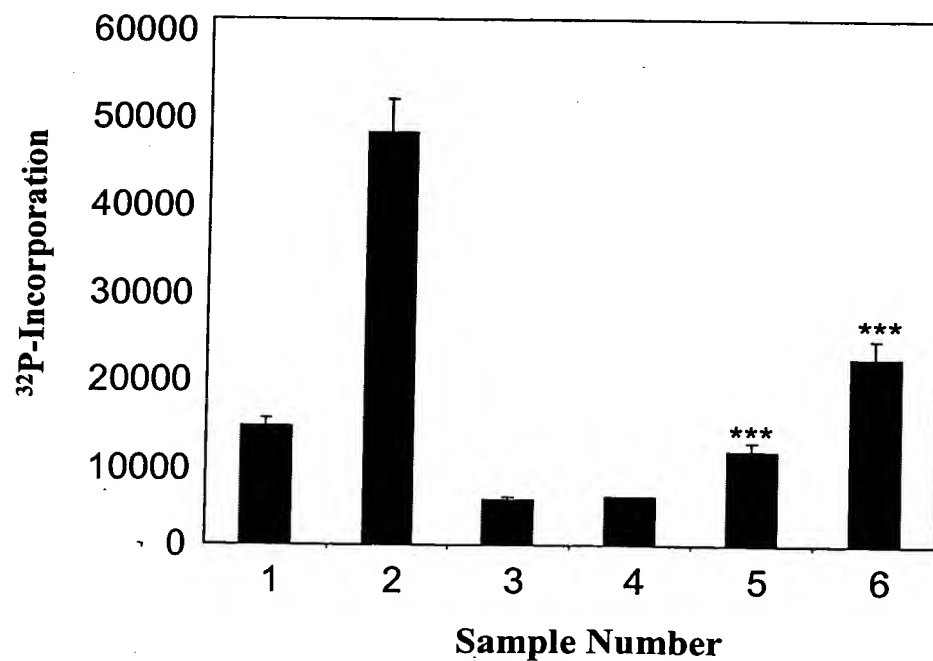
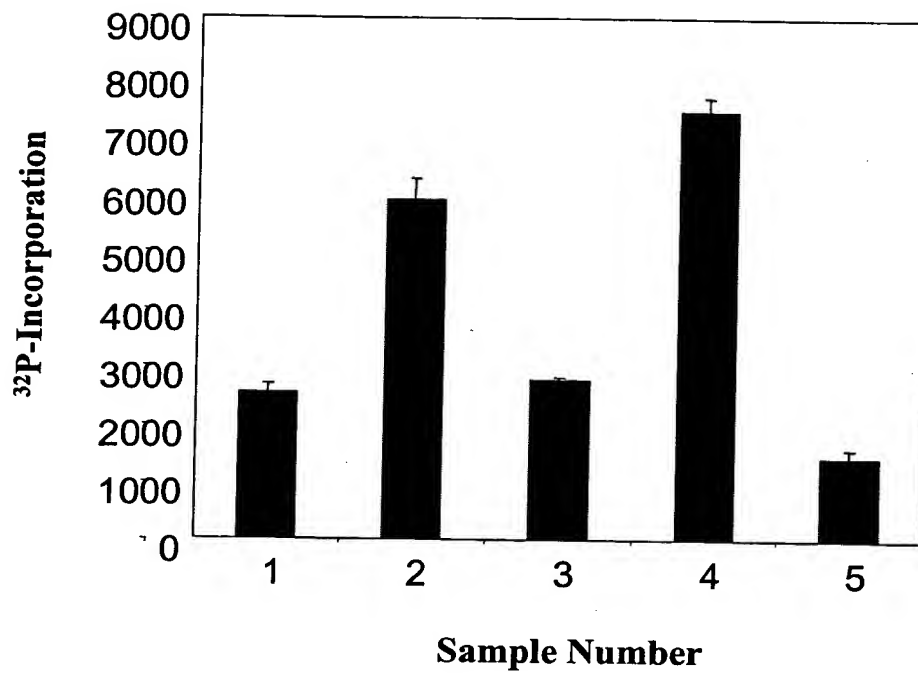
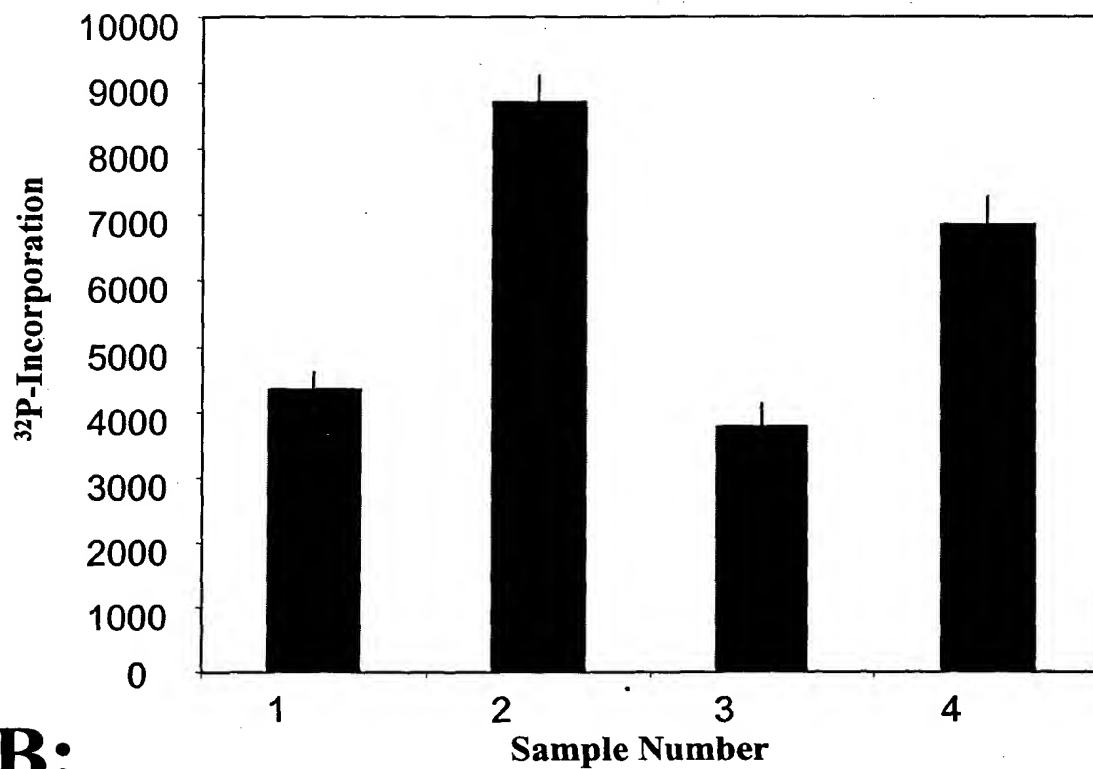
**B**

FIGURE 13

A:**B:****FIGURE 14**

**FIGURE 14 CONTINUED**

A:



B:

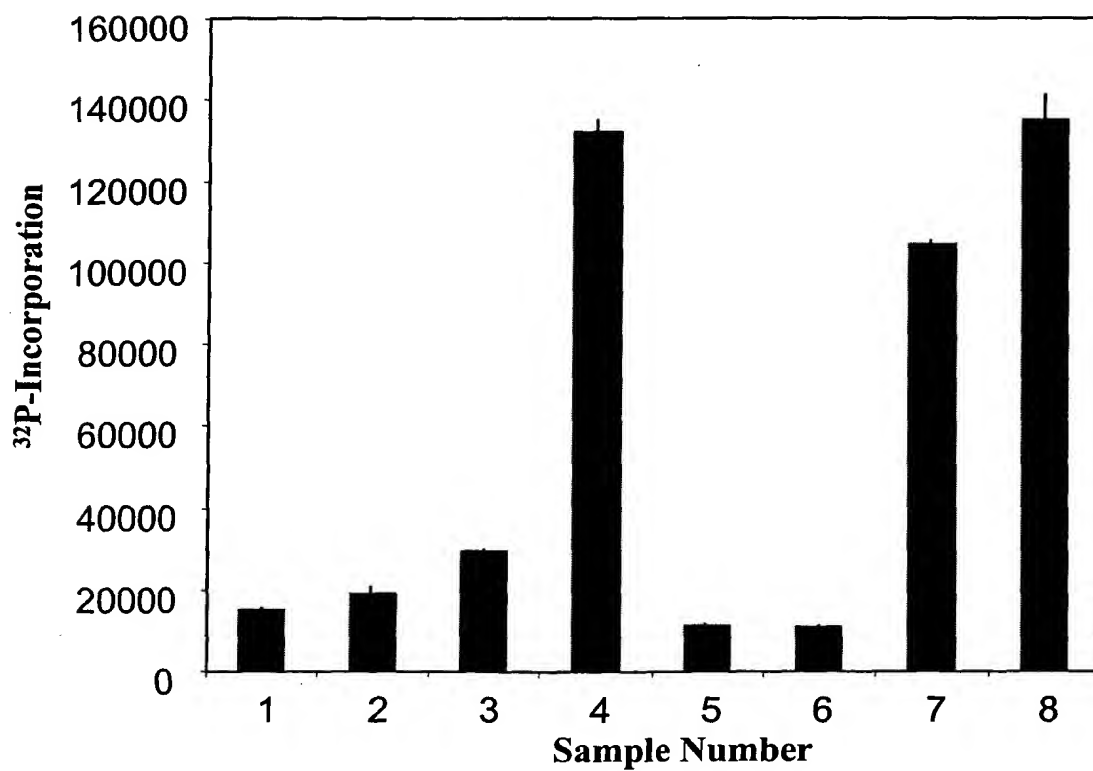


FIGURE 15

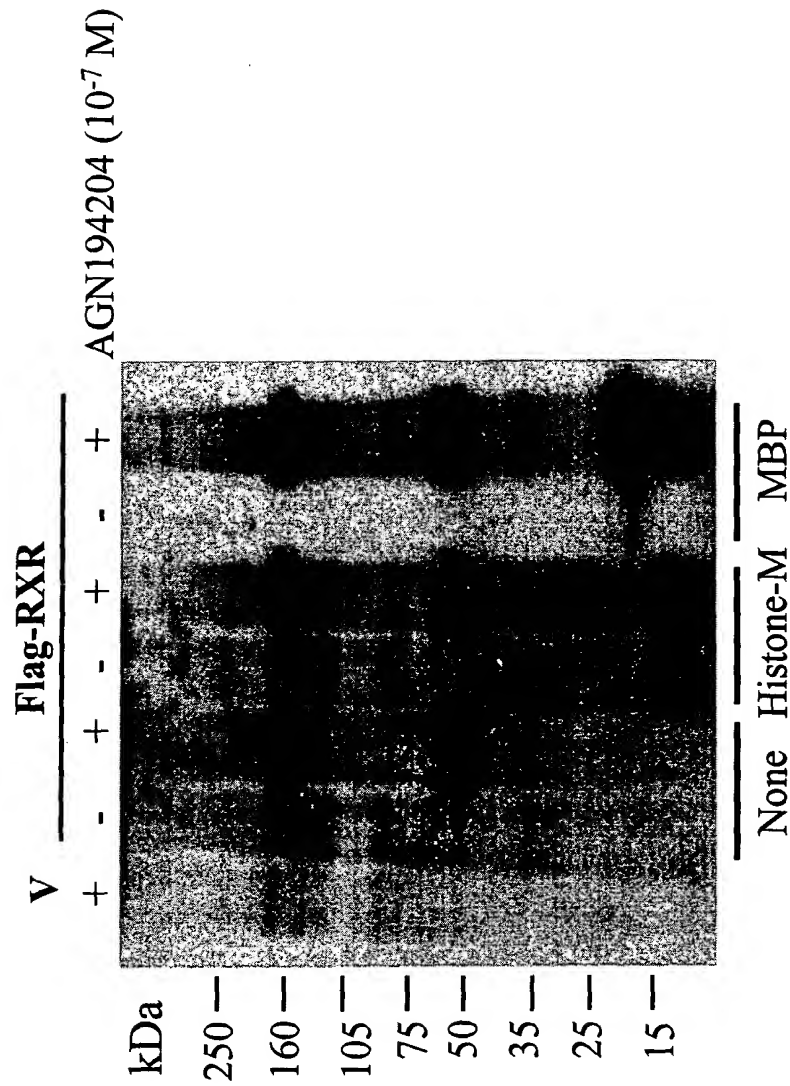
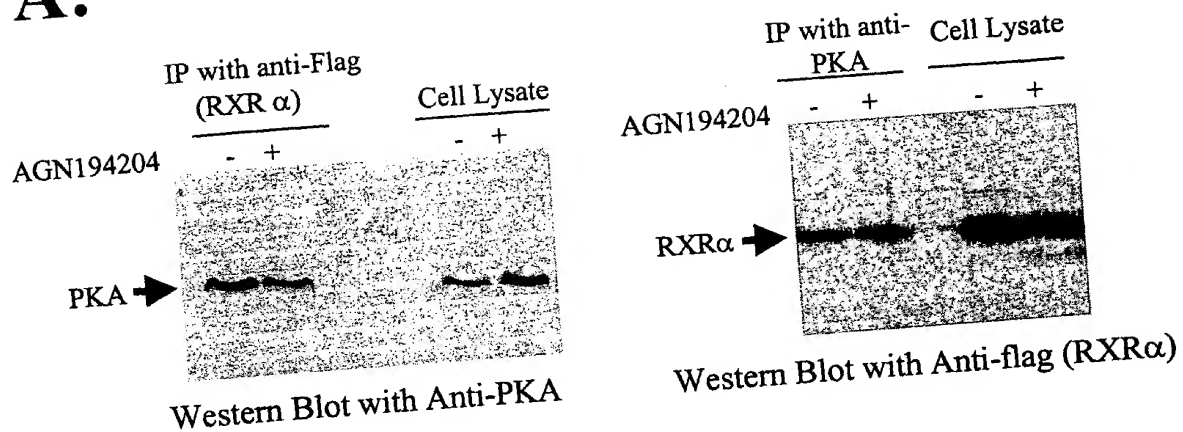


FIGURE 16

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A:



B:

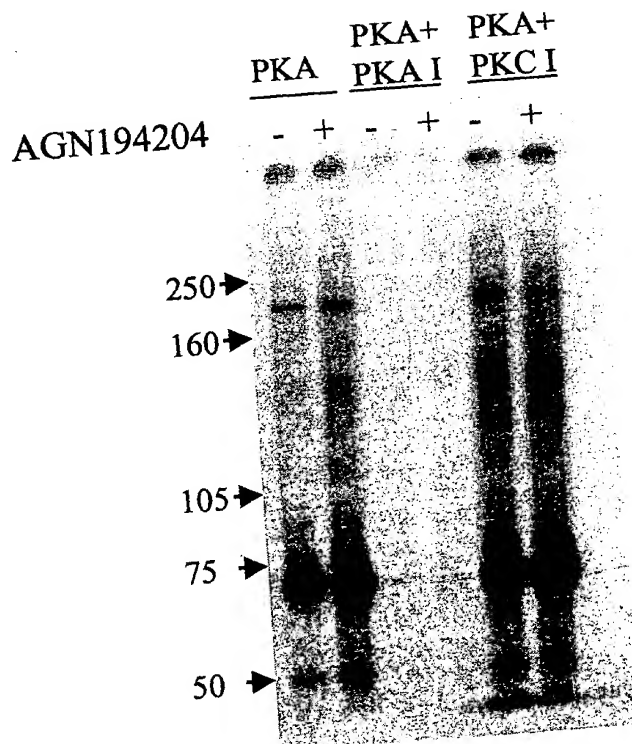


FIGURE 17

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Plec-N	GST	ISPCQD	FGKRMFVKIT	TTKQQDHFFQAAF	LEERDAMVRD	INKAIKICIEG
Plec-C	GCVVT	SVESN	SNGRK SEENL	FEII	TADEVHYFLOAAIT	IQMASRTGK*
RasGAP	VCSVY	VVHDS	LFGRPNCFOIV	VQHFS	...EE	HYIFYFAGET	PECAEDMYKG	IQAFCNLRKS
Akt	NFSVA	ECQLM	KTERPRPNTFVIR	CLQWT	...T	VIERTFHVDS	PDEREEMRA	IQMVANSLKQ
Spectrin	EAVCE	VALDY	KKKKHVFELR	LNDGNEYTFQAKD	DEEMNTWIOA	ISSATISSDKI
BARK	EQSVE	ETQI	KERKCLLLK	IRGGKQFIIQDCDS	DPELVQMKKE	LRCAVREAAQ
Tiam1-PH1	ENSTV	QAVPE	HPKKDFVFCLS	NSLGDAFLFQTTISQ	TELENW ITA	IHSACAAAVA
Grb7	ESNVY	VTQG	RKLYG MPTDFGFCVK	PNKLR	...NGH	KGLHIFCSED	EDSRTCMYAA	FRLFYGVQL
Dynammin	NLKL	RDVEKG	FMSSKHIFALF	NTEQRNVYKD	YROLELACET	QEEVDSMKAS	FLRAGVYPER	
Irs1	SCFN	INKRAD	SKNKHLVALY	TRDEHFALJAACS	EAEQDSMYQA	LLQLHNRKA
RasGRF-PH1	EGSIC	KRMP	PKRG	TSSKES	DKQHYFTW	FSNDSQKSLELRIDD	SKDCDEMVA	IARASYKILA
RasGRF-PH2	DCTI	LDPEN	MDDD	...GKGQ	EVDHLDKIM	VEPKD	...SP	PFTVILVASS	ROEKAAMTSD	IIQQVDNIR
Db1	EVGITE	YVKG	DNRKFEIM	YGEKEEVYIVQASN	VDKMTMDKE	IRNILLKQOE
Vav	SFQVR	DDSSG	ERDNKKWSHMFLLI	EDQGAQGVELFFKI	RELKKKIMEQ	FEMAISNIYP
Sos	KVQI	NDKDDT	NEYKHAFEII	LKDENSVIFSAKS	AEEKNNMMAA	LISLQYRSTL
Bcr	QIKSD	IQREK	RANK	<19>LL	MSPSMAFRVH	SRNGKSYTFLISS	DYERAEMREN	IREQQKKCFR
Btk	KITCV	ETVVP	EKNP	<20>II	ERFPYPFQW	YDEGPLYMFSPT	EEQKRMIHQ	LKNVIRYNSD
PLC-delta	DIQEV	RMGHR	TEGLE	KFARD	IPEDRCFSIV	FKDQRNTLDLIAPS	PRDAQHMVQG	LRKIITHSGS
PLC-gamma	EIKEI	RPGKT	SRDF	<7>AFR	PDQSHCFVIL	YGMF	...RL	KTLSLOATSE	DEVNMIIKG	ITWLMEDTLQ

FIG.18A

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Plec-N	..MEPKRIRE	GYLVKKGVS.FNT	WKPMWVILEDGI	EFYKKK.SD	NSPKGMIPLK
Plec-C	EEFRGVI IQ	GCLLKQHR.RKN	WKVRKFILREDPAYL	HYQPA.GA	EDPLGATHLR
RasGAP	DAFYKNIMKK	GYLLKKGKG.IKR	WKNLVFIILEGSDAQL	IYFESIKRA	TKPKGLICLS
Akt	.MNEVSVIKE	GMLHKRGEY.IKT	WRP YFQLKSDGSFI	GKERP.E	APDQTLPPIN
Spectrin	TQETPSAQME	GFLNRKHEWE	AHNKKASSRS	WHNVYCVINNQEM	GFYKDA.KTAASGIP	YHSEVPVSUK
BARK	YALGKDCIMH	GYMSKMGNP.FLTQ	WQRYFYILFPNRL	EWREG.EAPQSLLTME	
Tiam1-PH1	VRKAGALAVK	NFLVHKKNK.	.KVESATPRK	WKHYMWSLKGCTL	FFYETD.GRSRIDH	NSVPKHAVW
Grb7	LNAGSFPEIQ	GFLQLRGSG.	...RGSGRKL	WKRFCCFLRRSGL	YYSTKG.TSKDP	RHLQYVADVN
Dynamin	NQDEILVIRK	GMLTINNIG.	...IMGG	SKEYMFVITAENL	SWYKDD.EE	KEKAYMLSD
Irs1	TDGFSVVRKV	GYLRKPKS..	MHKRFVILRA	ASEAGGPRL	EYENE.KKWRHKS	SAPKRSIPLE
RasGRF-PH1	LLAQRDGTRK	GYLSKRSSD.NPK	WQTRWFAILLQNLL	FYQESD.S	SSRPSGIMLL
RasGRF-PH2	LDSQTFFVRQ	GSLIQVPMS	<10>GSLSLKK	EGEQCFILFSKHL	IICTRG.SGSKLH	LTKNGVLSLI
Db1	IGHKKGATKM	KDLARFK...	PMORHIFLYEKAT	VECKRR.	VE	SGEGSDRYP
Vav	LANYGRPKID	GELKITSVE.RRS	KTRVAFILDKAL	LICKRR.GDS	YDLKASVNLH
Sos	GQCCNEFTME	GILTRVGA..	KHERHIFLFDGL	MICCKSNHGQ	PRLPGASNAE	YRLKEKFFMR
Bcr	VFLFTDLLLC	TKLKQSGG.KTQQ	YDTCKWYPLTDL	SFQWIDELEA	VPNIPLVPDE	ELDALKIKLS
Btk	..MAAVILES	IFLKRQQK.	...KKTSPN	FKRQELLTVHKL	SWVEYD.FERGR	GSKKGSIDME
PLC-delta	DDPDLQALIK	GSQLLKVK.SSW	RRRFYKIQEDCK	TINGES.RKVMR	SPESQLFSLE
PLC-gamma	VLHLCRSLLEV	GTMTLFYS.	...KKSQPE	RKTFQVKLETROL	TISRGA.	DKIEGALDIR

FIG.18B